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Amendments to the Claims

This listing of claims replaces all prior versions and listings of claims in the application.

Listing of Claims

- 1. (Withdrawn) A method to reduce the sensitivity of endothelially-compromised vascular smooth muscle in a patient in need of such reduction, comprising administering a pharmaceutically effective amount of a CLC3 blocker, or a pharmaceutically acceptable salt thereof.
- 2. (Withdrawn) A method of claim 1, wherein the CLC3 blocker is a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6

wherein either R^4 is H or a lower alkyl radical and R^5 is a lower alkyl radical, or R^4 and R^5 are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

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3. (Withdrawn) A method of claim 2, wherein the compound administered is 1-p- β -dimethylaminoethoxyphenyl-trans-l, 2-diphenylbut-l-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.

- 4. (Withdrawn) A method to ameliorate the negative effects associated with vascular smooth muscle endothelium damage in a patient is need of such treatment, comprising administering a pharmaceutically effective amount of a CLC3 blocker, or a pharmaceutically acceptable salt thereof.
- 5. (Withdrawn) A method of claim 4, wherein the CLC3 blocker is a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6
 R^7

wherein either R^4 is H or a lower alkyl radical and R^5 is a lower alkyl radical, or R^4 and R^5 are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

or a pharmaceutically acceptable salt thereof.

6. (Withdrawn) A method of claim 5, wherein the wherein the compound administered is 1-p-β-dimethylaminoethoxyphenyl-trans-1,2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.

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7. (Withdrawn) A method of claim 5, wherein said endothelium damage is the result of diabetes.

- 8. (Withdrawn) A method of claim 5, wherein said endothelium damage is the result of a surgical procedure.
- 9. (Withdrawn) A method of claim 5, wherein said endothelium damage is the result or cause of hypertension.
- 10. (Withdrawn) A method of claim 5, wherein said endothelium damage is the result or cause of coronary artery disease.
- 11. (Withdrawn) A method of claim 5, which further comprises administering a pharmaceutically effective compound selected from the group consisting of: an anti-diabetes agent; anti-hypertension agent; an anti-coronary artery disease agent; and an anti-restensis agent.
- 12. (Withdrawn) A method to affect CLC3 receptors comprising administering a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6
 R^8

wherein either R^4 is H or a lower alkyl radical and R^5 is a lower alkyl radical, or R^4 and R^5 are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

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R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

or a pharmaceutically acceptable salt thereof.

- 13. (Withdrawn) A method of claim 12, wherein the compound administered is 1-p- β -dimethylaminoethoxyphenyl-trans-l, 2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.
- 14. (Withdrawn) A method to reduce contraction of endothelially-compromised vascular smooth muscle in response to agonist, comprising administering a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6

wherein either R⁴ is H or a lower alkyl radical and R⁵ is a lower alkyl radical, or R⁴ and R⁵ are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

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15. (Withdrawn) A method of claim 14, wherein the compound administered is l-p-β-dimethylaminoethoxyphenyl-trans-1,2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.

- 16. (Withdrawn) A method to decrease the effects of vasoconstrictors in pathologic tissues and not in non-pathologic tissues in a patient with pathologic tissues, and who is in need of such decrease, comprising administering a pharmaceutically-effective amount of a CLC3 blocker, or a pharmaceutically acceptable salt thereof.
- 17. (Withdrawn) A method of claim 16, wherein the CLC3 blocker is a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6
 R^7

wherein either R⁴ is H or a lower alkyl radical and R⁵ is a lower alkyl radical, or R⁴ and R⁵ are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

or a pharmaceutically acceptable salt thereof.

18. (Withdrawn) A method of claim 17, wherein the compound administered is l-p-β-dimethylaminoethoxyphenyl-trans-1,2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.

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19. (Withdrawn) A method to stabilize blood pressure in patients with endothelium compromised vascular smooth muscle, and who are in need of such stabilization, comprising administering a pharmaceutically-effective amount of a CLC3 blocker, or a pharmaceutically acceptable salt thereof.

20. (Withdrawn) A method of claim 19, wherein the CLC3 blocker is a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 C
 R^6

I

wherein either R^4 is H or a lower alkyl radical and R^5 is a lower alkyl radical, or R^4 and R^5 are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

- 21. (Withdrawn) A method of claim 20, wherein the compound administered is $l-p-\beta$ -dimethylaminoethoxyphenyl-trans-1,2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.
- 22. (Currently Amended) A method to modulate vascular tone in a patient having compromised vascular tissue, comprising administering a pharmaceutically effective amount of a chloride

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channel blocking agent, or a pharmaceutically acceptable salt thereof, wherein the compromised vascular tissue is associated with erectile dysfunction.

23. (Original) A method of claim 22, wherein the chloride channel blocking agent is a compound of Formula I

$$R^4R^5N(CH_2)_nO$$
 R^6
 R^8
 I

wherein either R⁴ is H or a lower alkyl radical and R⁵ is a lower alkyl radical, or R⁴ and R⁵ are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

- 24. (Currently Amended) A method of claim 23, wherein the wherein the compound is 1-p-β-dimethylaminoethoxyphenyl-trans-1,2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.
- 25. (Withdrawn) A method of claim 22, wherein the agent is selected from niflumic acid, mefanamic acid, flufenamic acid, 4,4'-diisothiocyanostilbene-2,2'-disulphonic acid (DIDS), 4,4'-dinitrostilbene-2,2'-disulphonic acid (DNDS), 4-acetamido-4'isiothiocyanostilbene-2,2'-disulphonic acid (SITS), anthracene-9-carboxylic acid (9-AC), 5-Nitro-2-(3-

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phenylpropylamino)benzoic acid (NPPB), diphenylamine-2-carboxylate (DPC), indanyloxyacetic acid-94 (IAA-94) and the pharmaceutically acceptable salts thereof.

- 26. (Withdrawn) The method of claim 25, wherein the agent is DIDS or a pharmaceutically acceptable salt thereof.
- 27. (Original) A method of claim 22, wherein the chloride channel is a CLC3 channel.
- 28. (Original) The method of claim 27, wherein blocking the CLC3 channel results in diminished vasoconstriction to norepinephrine.
- 29. (Original) The method of claim 22, wherein the agent modulates vascular tone by enhancing vasodilation.
- 30. (Canceled)
- 31. (Currently Amended) A method of claim 22, further comprising administering a pharmaceutically effective compound selected from an anti-diabetes agent, [[;]] an anti-hypertension agent, an anti-coronary artery disease agent, an anti-restenosis agent, and a vasodilatory agent.
- 32. (Original) A method of claim 22, wherein the agent is administered intravenously or orally.
- 33. (Currently Amended) A method to modulate penile vascular tone in a mammal <u>in need</u> thereof, said method comprising administering a pharmaceutically effective amount of a chloride channel blocking agent, or a pharmaceutically acceptable salt thereof.
- 34. (Original) A method of claim 33, wherein the chloride channel blocking agent is a compound of Formula I

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$$R^4R^5N(CH_2)_nO$$
 C
 R^6
 R^8

wherein either R⁴ is H or a lower alkyl radical and R⁵ is a lower alkyl radical, or R⁴ and R⁵ are joined together with the adjacent nitrogen atom to form a heterocyclic radical;

R⁶ is H or a lower alkyl radical;

R⁷ is H, halo, OH, a lower alkyl radical, or is a buta-1,3-dienyl radical which together with the adjacent benzene ring forms a naphthyl radical;

R⁸ is H or OH; and

n is 2;

- 35. (Currently Amended) A method of claim 34, wherein the compound administered is 1-p-β-dimethylaminoethoxyphenyl-trans-l, 2-diphenylbut-1-ene (tamoxifen), or a pharmaceutically acceptable salt thereof.
- 36. (Withdrawn) A method of claim 33, wherein the agent is selected from niflumic acid, mefanamic acid, flufenamic acid, 4,4'-diisothiocyanostilbene-2,2'-disulphonic acid (DIDS), 4,4'-dinitrostilbene-2,2'-disulphonic acid (DNDS), 4-acetamido-4'isiothiocyanostilbene-2,2'-disulphonic acid (SITS), anthracene-9-carboxylic acid (9-AC), 5-Nitro-2-(3-phenylpropylamino)benzoic acid (NPPB), diphenylamine-2-carboxylate (DPC), indanyloxyacetic acid-94 (IAA-94) and the pharmaceutically acceptable salts thereof.
- 37. (Withdrawn) The method of claim 36, wherein the agent is DIDS or a pharmaceutically acceptable salt thereof.

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38. (Original) The method of claim 33, wherein the agent is administered orally or intravenously.

- 39. (Original) A method of claim 33, wherein the chloride channel is a CLC3 channel.
- 40. (Original) The method of claim 39, wherein blocking the CLC3 channel results in diminished vasoconstriction to norepinephrine.
- 41. (Original) The method of claim 39, wherein blocking the CLC3 channel reduces penile sympathetic tone.
- 42. (Original) The method of claim 41, wherein the reduction of penile sympathetic tone induces an erection.
- 43. (Withdrawn) A method for treating erectile dysfunction comprising administering a composition comprising a CLC3 channel blocking agent or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.